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175136-87-5/RN
E1
E2
                    175136-88-6/RN
E3
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                    175136-90-0/RN
E4
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                    175136-91-1/RN
              1
E5
                    175136-92-2/RN
E6
              1
                    175136-93-3/RN
E7
              1
                    175136-94-4/RN
E8
              1
                    175136-95-5/RN
E9
              1
                    175136-96-6/RN
              1
E10
                    175136-97-7/RN
E11
              1
E12
                    175136-98-8/RN
              1
=> s e3
             1 175136-89-7/RN
L1
=> d l1
```

L1 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS

RN 175136-89-7 REGISTRY

CN Benzenemethanamine, 2-chloro-6-phenoxy- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 2-Aminomethyl-3-chlorodiphenyl ether

CN 2-Chloro-6-phenoxybenzylamine

FS 3D CONCORD

MF C13 H12 Cl N O

SR CAS Registry Services

LC STN Files: CA, CAPLUS, CHEMCATS, CSCHEM, USPATZ, USPATFULL

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3 REFERENCES IN FILE CA (1962 TO DATE)

3 REFERENCES IN FILE CAPLUS (1962 TO DATE)

=> FIL REGISTRY

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 2.08 2.29

FILE 'REGISTRY' ENTERED AT 16:25:19 ON 01 APR 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

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STRUCTURE FILE UPDATES: 31 MAR 2003 HIGHEST RN 501072-24-8 DICTIONARY FILE UPDATES: 31 MAR 2003 HIGHEST RN 501072-24-8

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

=> logoff

ALL L# QUERIES AND ANSWER SETS ARE DELETED AT LOGOFF

LOGOFF? (Y)/N/HOLD:Y

COST IN U.S. DOLLARS

SINCE FILE TOTAL
ENTRY SESSION

FULL ESTIMATED COST 46.19 194.55

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION

CA SUBSCRIBER PRICE -6.51 -6.51

STN INTERNATIONAL LOGOFF AT 09:05:22 ON 04 FEB 2003

NEWS 39

NEWS 40

NEWS 42 Jan 29

Jan 13

Connecting via Winsock to STN

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Welcome to STN International! Enter x:X
LOGINID:ssspta1626kas
PASSWORD:
TERMINAL (ENTER 1, 2, 3, OR ?):2
                     Welcome to STN International
                 Web Page URLs for STN Seminar Schedule - N. America
NEWS
NEWS 2 Apr 08 "Ask CAS" for self-help around the clock
NEWS 3 Apr 09 BEILSTEIN: Reload and Implementation of a New Subject Area
NEWS 4 Apr 09 ZDB will be removed from STN
NEWS 5 Apr 19 US Patent Applications available in IFICDB, IFIPAT, and IFIUDB
NEWS 6 Apr 22 Records from IP.com available in CAPLUS, HCAPLUS, and ZCAPLUS
NEWS 7 Apr 22 BIOSIS Gene Names now available in TOXCENTER
NEWS 8 Apr 22 Federal Research in Progress (FEDRIP) now available
NEWS 9 Jun 03 New e-mail delivery for search results now available
NEWS 10 Jun 10 MEDLINE Reload
NEWS 11 Jun 10 PCTFULL has been reloaded
NEWS 12 Jul 02 FOREGE no longer contains STANDARDS file segment
         Jul 22 USAN to be reloaded July 28, 2002;
NEWS 13
                 saved answer sets no longer valid
NEWS 14 Jul 29 Enhanced polymer searching in REGISTRY
NEWS 15 Jul 30 NETFIRST to be removed from STN
NEWS 16 Aug 08 CANCERLIT reload
NEWS 17 Aug 08 PHARMAMarketLetter(PHARMAML) - new on STN
NEWS 18 Aug 08 NTIS has been reloaded and enhanced
                 Aquatic Toxicity Information Retrieval (AQUIRE)
NEWS 19
         Aug 19
                 now available on STN
         Aug 19 IFIPAT, IFICDB, and IFIUDB have been reloaded
NEWS 20
        Aug 19 The MEDLINE file segment of TOXCENTER has been reloaded
NEWS 21
                 Sequence searching in REGISTRY enhanced
NEWS 22
         Aug 26
                 JAPIO has been reloaded and enhanced
NEWS 23
         Sep 03
NEWS 24 Sep 16 Experimental properties added to the REGISTRY file
NEWS 25 Sep 16 CA Section Thesaurus available in CAPLUS and CA
NEWS 26 Oct 01 CASREACT Enriched with Reactions from 1907 to 1985
NEWS 27 Oct 21 EVENTLINE has been reloaded
NEWS 28 Oct 24 BEILSTEIN adds new search fields
NEWS 29 Oct 24 Nutraceuticals International (NUTRACEUT) now available on STN
NEWS 30 Oct 25 MEDLINE SDI run of October 8, 2002
NEWS 31 Nov 18 DKILIT has been renamed APOLLIT
NEWS 32 Nov 25 More calculated properties added to REGISTRY
NEWS 33 Dec 02 TIBKAT will be removed from STN
NEWS 34 Dec 04 CSA files on STN
NEWS 35 Dec 17 PCTFULL now covers WP/PCT Applications from 1978 to date
NEWS 36 Dec 17
                TOXCENTER enhanced with additional content
                Adis Clinical Trials Insight now available on STN
NEWS 37
         Dec 17
        Dec 30 ISMEC no longer available
NEWS 38
```

Jan 21 NUTRACEUT offering one free connect hour in February 2003

NEWS 41 Jan 21 PHARMAML offering one free connect hour in February 2003

Indexing added to some pre-1967 records in CA/CAPLUS

Simultaneous left and right truncation added to COMPENDEX,

ENERGY, INSPEC

NEWS EXPRESS
January 6 CURRENT WINDOWS VERSION IS V6.01a,
CURRENT MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP),
AND CURRENT DISCOVER FILE IS DATED 01 OCTOBER 2002
NEWS HOURS
STN Operating Hours Plus Help Desk Availability
NEWS INTER
General Internet Information
NEWS LOGIN
NEWS PHONE
Direct Dial and Telecommunication Network Access to STN
NEWS WWW
CAS World Wide Web Site (general information)

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FILE 'HOME' ENTERED AT 09:03:32 ON 04 FEB 2003

=> file reg
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

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STRUCTURE FILE UPDATES: 2 FEB 2003 HIGHEST RN 484639-64-7 DICTIONARY FILE UPDATES: 2 FEB 2003 HIGHEST RN 484639-64-7

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> Testing the current file.... screen

ENTER SCREEN EXPRESSION OR (END):end

=>

Uploading C:\Program Files\Stnexp\Queries\0984599211.str

L1 STRUCTURE UPLOADED

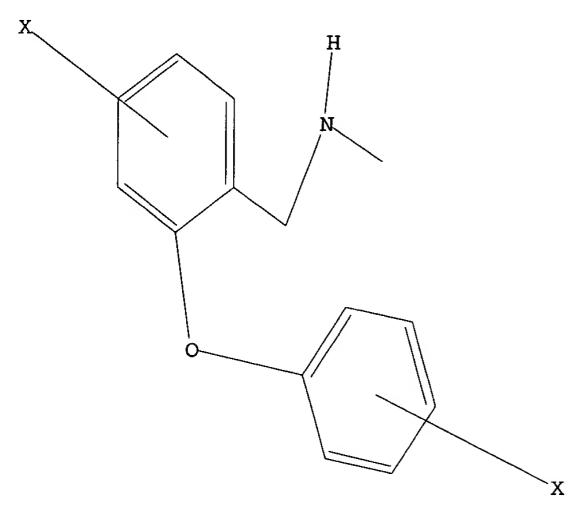
=> que L1

L2 QUE L1

=> d

L2 HAS NO ANSWERS

L1 STR



Structure attributes must be viewed using STN Express query preparation. L2 QUE ABB=ON PLU=ON L1

=> s l1 full

FULL SEARCH INITIATED 09:04:16 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 2182 TO ITERATE

100.0% PROCESSED 2182 ITERATIONS

33 ANSWERS

148.36

SEARCH TIME: 00.00.01

L3 33 SEA SSS FUL L1

=> file caplus

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION

148.15

FULL ESTIMATED COST

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FILE COVERS 1907 - 4 Feb 2003 VOL 138 ISS 6 FILE LAST UPDATED: 3 Feb 2003 (20030203/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 13 L4 10 L3

=> d ibib abs hitstr tot

L4 ANSWER 1 OF 10 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2002:925264 CAPLUS DOCUMENT NUMBER: 138:11431 TITLE: 5-HT1a antagonist or an .alpha.2-adrenergic antagonist in combination with an serotonin reuptake inhibitor for treatment of sleep disorders, including sleep apnea Howard, Harry Ralph, Jr. INVENTOR (S): PATENT ASSIGNEE(S): Pfizer Products Inc., USA SOURCE: Eur. Pat. Appl., 22 pp. CODEN: EPXXDW DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE EP 1262197 A2 20021204 EP 2002-253589 20020522 EP 1262197 A3 20021218 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR US 2002183306 A1 20021205 US 2002-75849 20020213 JP 2003026602 A2 20030129 JP 2002-155222 20020529 PRIORITY APPLN. INFO.: US 2001-294322P P 20010530 OTHER SOURCE(S): MARPAT 138:11431 The invention provides a method of treating sleep disorders, including sleep apnea, in a mammal, including a human, by administering to the mammal a 5-HT1a antagonist or an .alpha.2-adrenergic antagonist in combination with an serotonin reuptake inhibitor (SRI) antidepressant agent with improvement in efficacy. Also provided are pharmaceutical compns. contg. a pharmaceutically acceptable carrier, a 5-HT1a antagonist or an .alpha.2-adrenergic antagonist, and an SRI antidepressant agent. 289716-94-5 289717-18-6 289717-57-3 289717-60-8 289717-61-9 289717-67-5 289717-68-6 289717-71-1 289717-72-2 444888-21-5 444888-23-7 444888-24-8 444888-49-7 477337-55-6 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (5-HT1a antagonist or .alpha.2-adrenergic antagonist in combination with serotonin reuptake inhibitor for treatment of sleep disorders, including sleep apnea) 289716-94-5 CAPLUS

L4 ANSWER 1 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)
CN Benzenemethanamine, 5-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA INDEX NAME)

Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA

Br C1

RN 289717-67-5 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-,
(+)- (9CI) (CA INDEX NAME)

Rotation (+).

Me NHMe C1

RN 289717-68-6 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-,
(-)- (9CI) (CA INDEX NAME)

Rotation (-).

Me NHMe C1

RN 289717-71-1 CAPLUS
CN Benzenemethanamine, 2-(4-chloro-3-fluorophenoxy)-5-fluoro-N-methyl- (9CI)
(CA INDEX NAME)

L4 ANSWER 1 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

MeNH-CH₂

RN 289717-18-6 CAPLUS
CN Benzenemethanamine, 4-chloro-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA INDEX NAME)

MeNH-CH₂

RN 289717-57-3 CAPLUS
CN Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA INDEX NAME)

Menh-CH₂

RN 289717-60-8 CAPLUS
CN Benzenemethanamine, 4-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9C1) (CA INDEX NAME)

MeNH-CH2

O

C1

RN 289717-61-9 CAPLUS

L4 ANSWER 1 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

MeNH-CH2

RN 289717-72-2 CAPLUS
CN Benzenemethanamine, 2-(3-chloro-4-fluorophenoxy)-5-fluoro-N-methyl- (9CI)
(CA INDEX NAME)

MeNH—CH₂

RN 444888-21-5 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.,.alpha.trimethyl- (9CI) (CA INDEX NAME)

NHMe Me

N 444888-23-7 CAPLUS N Benzenemethanamine, 4-chloro-2-(4-chlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA INDEX NAME)

MeNH-CH2

ANSWER 1 OF 10 CAPLUS COPYRIGHT 2003 ACS L4(Continued) 444888-24-8 CAPLUS RN

Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-4-methoxy-N-methyl-(9CI) (CA INDEX NAME)

MeNH-CH2

444888-49-7 CAPLUS

Benzenemethanamine, 4-chloro-2-(3,4-dichlorophenoxy)-N-methyl-5-

(methylsulfonyl) - (9CI) (CA INDEX NAME)

477337-55-6 CAPLUS Benzenemethanamine, 5-fluoro-2-(4-fluoro-3-methoxyphenoxy)-N-methyl-

(9CI) (CA INDEX NAME)

ANSWER 2 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

for use in alcoholism and alc. dependence) 289716-94-5 CAPLUS

Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA INDEX NAME)

289717-18-6 CAPLUS Benzenemethanamine, 4-chloro-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA

CN INDEX NAME)

289717-24-4 CAPLUS

Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-

(9CI) (CA INDEX NAME)

289717-57-3 CAPLUS

Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA INDEX NAME)

L4 ANSWER 2 OF 10 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2002:925263 CAPLUS

DOCUMENT NUMBER: 138:336

Combination of a monoamine reuptake inhibitor and an TITLE: opioid antagonist for use in alcoholism and alcohol

dependence

INVENTOR (S): Howard, Harry Ralph, Jr. PATENT ASSIGNEE(S): Pfizer Products Inc., USA SOURCE:

Eur. Pat. Appl., 37 pp. CODEN: EPXXDW

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

> PATENT NO. KIND DATE APPLICATION NO. DATE EP 1262196 20021204 EP 2002-253105 20020502 EP 1262196 A3 20021218

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR JP 2002370975 A2 20021224 JP 2002-132804 20020508

AU 2002040686 A5 20021205 AU 2002-40686 20020516 CN 1386503 A 20021225 CN 2002-120350 20020523 PRIORITY APPLN. INFO.: US 2001-293088P P 20010523

OTHER SOURCE(S): MARPAT 138:336

The present invention relates to a method of treating alcoholism or alc. dependence in a mammal, including a human, by administering to the mammal a monoamine reuptake inhibitor in combination with an opioid antagonist. It also relates to pharmaceutical compns. contg. a pharmaceutically

acceptable carrier, a monoamine reuptake inhibitor and an opioid antagonist. An example monoamine reuptake inhibitor is I.

289716-94-5 289717-18-6 289717-24-4 289717-57-3 289717-58-4 289717-60-8

289717-61-9 289717-67-5 289717-68-6 289717-71-1 289717-72-2

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (combination of a monoamine reuptake inhibitor and an opioid

ANSWER 2 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

MeNH-

289717-58-4 CAPLUS Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-(9CI)

(CA INDEX NAME)

289717-60-8 CAPLUS RN

Benzenemethanamine, 4-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA ÇN

менн-сн2

289717-61-9 CAPLUS

Benzenemethanamine, 5-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA

MeNH-CH2

289717-67-5 CAPLUS

Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-, (+) - (9CI) (CA INDEX NAME)

L4 ANSWER 2 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued) Rotation (+).

Me NHMe C1

RN 289717-68-6 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-,
(-)- (9CI) (CA INDEX NAME)

Rotation (-).

Me_NHMe

RN 289717-71-1 CAPLUS
CN Benzenemethanamine, 2-(4-chloro-3-fluorophenoxy)-5-fluoro-N-methyl- (9CI)
(CA INDEX NAME)

MeNH-CH₂
C1

ACCESSION NUMBER:

DOCUMENT NUMBER:

TITLE:

RN 289717-72-2 CAPLUS
CN Benzenemethanamine, 2-(3-chloro-4-fluorophenoxy)-5-fluoro-N-methyl- (9CI)
(CA INDEX NAME)

L4 ANSWER 2 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

MeNH-CH₂

inhibitor and dopamine D3 receptor agonist for depression and anxiety INVENTOR(S): Howard, Harry Ralph, Jr PATENT ASSIGNEE(S): Pfizer Products Inc., USA SOURCE: Eur. Pat. Appl., 31 pp. CODEN: EPXXDW DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE EP 1260221 20021127 EP 2002-253135 20020503 EP 1260221 A3 20021218 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR AU 2002040681 A5 20021205 AU 2002-40681 20020516

2002:904325 CAPLUS

Combination treatment with monoamine reuptake

L4 ANSWER 3 OF 10 CAPLUS COPYRIGHT 2003 ACS

JP 2002-141515 JP 2002370976 A2 20021224 20020516 CN 1386504 20021225 CN 2002-120351 20020523 PRIORITY APPLN. INFO.: US 2001-293063P P 20010523 OTHER SOURCE(S): MARPAT 137:380038 The present invention relates to a method of treating depression or anxiety in a mammal, including a human, by administering to the mammal a monoamine reuptake inhibitor in combination with a dopamine D3 receptor agonist. It also relates to pharmaceutical compns. contg. a pharmaceutically acceptable carrier, a monoamine reuptake inhibitor and a dopamine D3 receptor agonist. 289716-94-5 289717-18-6 289717-24-4 289717-57-3 289717-58-4 289717-60-8 289717-61-9 289717-67-5 289717-68-6 289717-71-1 289717-72-2

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (monoamine reuptake inhibitor; combination treatment with monoamine reuptake inhibitor and dopamine D3 receptor agonist for depression and anxiety)
289716-94-5 CAPLUS
Benzenemethanamine, 2-{3,4-dichlorophenoxy}-5-fluoro-N-methyl- (9CI) (CA

MeNH-CH₂

INDEX NAME)

RN 289717-18-6 CAPLUS
CN Benzenemethanamine, 4-chloro-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA INDEX NAME)

L4 ANSWER 3 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

MeNH-CH₂

RN 289717-24-4 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl(9CI) (CA INDEX NAME)

NHMe CH-Me

RN 289717-57-3 CAPLUS
CN Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA INDEX NAME)

MeNH-CH2

RN 289717-58-4 CAPLUS
CN Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N,.alpha.-dimethyl(9CI)
(CA INDEX NAME)

NHMe CH- Me

RN 289717-60-8 CAPLUS
CN Benzenemethanamine, 4-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA

ANSWER 3 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued) INDEX NAME)

MeNHT CH2

289717-61-9 CAPLUS Benzenemethanamine, 5-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA

MeNH-

289717-67-5 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-, (+) - (9CI) (CA INDEX NAME)

Rotation (+).

289717-68-6 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-, (-) - (9CI) (CA INDEX NAME)

Rotation (-).

ANSWER 3 OF 10 CAPLUS COPYRIGHT 2003 ACS

289717-71-1 CAPLUS Benzenemethanamine, 2-(4-chloro-3-fluorophenoxy)-5-fluoro-N-methyl- (9CI) CN (CA INDEX NAME)

MeNH-CH2

289717-72-2 CAPLUS Benzenemethanamine, 2-(3-chloro-4-fluorophenoxy)-5-fluoro-N-methyl- (9CI) (CA INDEX NAME)

L4 ANSWER 4 OF 10 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2002:755211 CAPLUS DOCUMENT NUMBER: TITLE: Preparation of phenoxybenzylamines as monoamine reuptake inhibitors for treatment of CNS disorders. INVENTOR (S): Howard, Harry R.; Schmidt, Christopher J.; Seeger, Thomas F.; Elliott, Mark L. PATENT ASSIGNEE (S): U.S. Pat. Appl. Publ., 24 pp., Cont.-in-part of U.S. SOURCE: Ser. No. 529,207. CODEN: USXXCO

Patent

English

FAMILY ACC. NUM. COUNT:

DOCUMENT TYPE:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE US 2002143003 A1 20021003 US 2001-845992 20010430 WO 2000050380 A1 20000831 WO 2000-IB108 20000202 W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG PRIORITY APPLN. INFO.: US 1999-121313P P 19990223 US 2000-529207 A2 20000202 WO 2000-IB108 W 20000202

OTHER SOURCE(S): MARPAT 137:262839 ROZCR3R4NR1R2 [R = (un)substituted Ph; R1,R2 = H, alk(en)yl, alkynyl; NR1R2 = heterocyclyl; R3,R4 = H or (fluoro)alkyl; R3R4 = (un)substituted alkylene; R2R3 = atoms to complete a heterocyclic ring; Z = (un) substituted phenylene) were prepd. as monoamine reuptake inhibitors (no data). Such compds. are useful exhibit activity as serotonin, norepinephrine and dopamine reuptake inhibitors, and their pharmaceutically acceptable salts, and their use in the treatment of central nervous system and other disorders.

289716-89-8P, Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-Nmethyl-, hydrochloride 289716-93-4P, Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N-methyl-, hydrochloride 289716-94-5P, Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N-methyl- 289716-95-6P, Benzenemethanamine, 2-(3,4dichlorophenoxy)-5-fluoro-N-methyl-, (2Z)-2-butenedioate (2:1) 289717-18-6P, Benzenemethanamine, 4-chloro-2-(3,4-dichlorophenoxy)-N-methyl- 289717-19-79, Benzenemethanamine, 4-chloro-2-(3,4dichlorophenoxy) -N-methyl-, (2Z)-2-butenedicate (1:1) 289717-23-3P , Benzenemethanamine, 2-(3,4-dichlorophenoxy)-3-fluoro-N-methyl-, (2Z)-2-butenedioate (1:1) 289717-24-4P, Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl- 289717-25-5P Benzenemethanamine,

2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-, (2Z)-2-butenedioate (1:1) 289717-26-6P, Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-, hydrochloride 289717-28-8P, Benzenemethanamine, 2-(3,4-dichlorophenoxy)-4-fluoro-N-methyl-, (2Z)-2-butenedicate (1:1) 289717-35-7P, Benzenemethanamine, 5-bromo-2-(3,4-dichlorophenoxy)-N-methyl-,

ANSWER 4 OF 10 CAPLUS COPYRIGHT 2003 ACS hydrochloride 289717-36-8P, Benzenemethanamine, 4-bromo-2-(3,4-dichlorophenoxy)-N-methyl-, hydrochloride 289717-57-3P, Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-Nmethyl- 289717-58-4P, Benzenemethanamine, 2-(4-chlorophenoxy)-5fluoro-N, .alpha.-dimethyl- 289717-60-8P, Benzenemethanamine, 4-bromo-2-(3,4-dichlorophenoxy)-N-methyl- 289717-61-9P, Benzenemethanamine, 5-bromo-2-(3,4-dichlorophenoxy)-N-methyl-289717-67-5P, Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N, alpha. dimethyl-, (+) - 289717-68-6P, Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N, .alpha.-dimethyl-, (-)-289717-71-1P, Benzenemethanamine, 2-(4-chloro-3-fluorophenoxy)-5fluoro-N-methyl- 289717-72-2P, Benzenemethanamine, 2-(3-chloro-4-fluorophenoxy)-5-fluoro-N-methyl-RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (prepn. of phenoxybenzylamines as monoamine reuptake inhibitors) 289716-89-8 CAPLUS Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N-methyl-, hydrochloride (9CI) (CA INDEX NAME)

MeNH-CH2

● HCl

289716-93-4 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N-methyl-, hydrochloride (9CI) (CA INDEX NAME)

MeNH-CH2

289716-94-5 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA L4 ANSWER 4 OP 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

MeNH-CH2

289716-95-6 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N-methyl-, (22)-2-butenedioate (2:1) (9CI) (CA INDEX NAME)

CM 1

CRN 289716-94-5 CMF C14 H12 C12 F N O

меин-сн2

CM 2

CRN 110-16-7 CMF C4 H4 O4

Double bond geometry as shown.

со2н

RN 289717-18-6 CAPLUS Benzenemethanamine, 4-chloro-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA INDEX NAME)

L4 ANSWER 4 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

MeNH-CH2

CRN 110-16-7

CMF C4 H4 Q4

Double bond geometry as shown.

CO₂H

289717-24-4 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-(9CI) (CA INDEX NAME)

289717-25-5 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CRN 289717-24-4

CMF C15 H14 C12 F N O

L4 ANSWER 4 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

менн-сн2

289717-19-7 CAPLUS Benzenemethanamine, 4-chloro-2-(3,4-dichlorophenoxy)-N-methyl-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 289717-18-6

CMF C14 H12 C13 N O

Мейн- Сн2

CM

CRN 110-16-7 CMF C4 H4 O4

Double bond geometry as shown.

CO2H

RN 289717-23-3 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-3-fluoro-N-methyl-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 289717-22-2 CMF C14 H12 C12 F N O

L4 ANSWER 4 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

ŅНМе CH- Me

CM 2

CRN 110-16-7 CMF C4 H4 O4

Double bond geometry as shown.

RN 289717-26-6 CAPLUS

Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-, hydrochloride (9CI) (CA INDEX NAME)

HC1

289717-28-8 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-4-fluoro-N-methyl-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CRN 289717-27-7

CMF C14 H12 C12 F N O

L4 ANSWER 4 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

MeNH-CH₂

CM 2

CRN 110-16-7 CMF C4 H4 O4

Double bond geometry as shown.

но₂с Z

RN 289717-35-7 CAPLUS
CN Benzenemethanamine, 5-bromo-2-(3,4-dichlorophenoxy)-N-methyl-,
hydrochloride (9CI) (CA INDEX NAME)

Br Cl

● HCl

RN 289717-36-8 CAPLUS
CN Benzenemethanamine, 4-bromo-2-(3,4-dichlorophenoxy)-N-methyl-,
hydrochloride (9CI) (CA INDEX NAME)

ANSWER 4 OF 10 CAPLUS COPYRIGHT 2003 ACS

MeNH-CH₂

RN 289717-61-9 CAPLUS
CN Benzenemethanamine, 5-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA INDEX NAME)

(Continued)

MeNH-CH2

RN 289717-67-5 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-,
(+)- (9CI) (CA INDEX NAME)

Rotation (+).

Me NHMe C1

RN 289717-68-6 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-,
(-)- (9CI) (CA INDEX NAME)

Rotation (-).

L4 ANSWER 4 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

MeNH-CH₂
Cl

● HCl

RN 289717-57-3 CAPLUS
CN Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA INDEX NAME)

MeNH—CH₂

RN 289717-58-4 CAPLUS
CN Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N,.alpha.-dimethyl(9CI)
(CA INDEX NAME)

NHMe CH—Me

RN 289717-60-8 CAPLUS
CN Benzenemethanamine, 4-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA INDEX NAME)

L4 ANSWER 4 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

Me NHMe

RN 289717-71-1 CAPLUS
CN Benzenemethanamine, 2-(4-chloro-3-fluorophenoxy)-5-fluoro-N-methyl- (9CI)
(CA INDEX NAME)

MeNH-CH2

MeNH-CH₂

IT 289718-10-1P, Benzamide, 2-(3.4-dichlorophenoxy)-5-fluoro-N-methyl-RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (prepn. of phenoxybenzylamines as monoamine reuptake inhibitors)

RN 289718-10-1 CAPLUS
CN Benzamide, 2-(3,4-dichlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA INDEX NAME)

L4 ANSWER 4 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

DOCUMENT NUMBER: 137:195595 TITLE: Atypical antipsychotic-antidepressant combination for treatment of depression, obsessive compulsive disorder, and psychosis Howard, Harry R., Jr. INVENTOR(S): Pfizer Inc., USA PATENT ASSIGNEE(S): SOURCE: U.S. Pat. Appl. Publ., 20 pp. CODEN: USXXCO DOCUMENT TYPE: Patent. LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE ------US 2002123490 A1 20020905 US 2001-10651 20011206 EP 2002-251153 20020220 EP 1238676 A1 20020911 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR JP 2002308801 A2 20021023 JP 2002-50579 20020227 PRIORITY APPLN. INFO.: US 2001-272619P P 20010301 OTHER SOURCE(S): MARPAT 137:195595 AB The invention provides a method for treating depression, obsessive compulsive disorder, and psychosis in a mammal, including a human, by administering to the mammal an atypical antipsychotic in combination with an antidepressant agent with improvement in efficiency. It also provides pharmaceutical compns. contg. a pharmaceutically acceptable carrier, an atypical antipsychotic, and a serotonin reuptake inhibitor. 289716-94-5 289717-18-6 289717-24-4 289717-57-3 289717-58-4 289717-60-8 289717-61-9 289717-67-5 289717-68-6 289717-71-1 289717-72-2 444888-21-5 444888-23-7 444888-24-8 444888-30-6 444888-49-7 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (atypical antipsychotic-antidepressant combination for treatment of depression, obsessive compulsive disorder, and psychosis) 289716-94-5 CAPLUS CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA

2002:674788 CAPLUS

L4 ANSWER 5 OF 10 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER:

MeNH-CH₂

INDEX NAME)

RN 289717-18-6 CAPLUS

L4 ANSWER 5 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)
CN Benzenemethanamine, 4-chloro-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA INDEX NAME)

MeNH-CH₂
Cl Cl

RN 289717-24-4 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl(9CI) (CA INDEX NAME)

NHMe CH-Me

RN 289717-57-3 CAPLUS
CN Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA INDEX NAME)

MeNH-CH2

RN 289717-58-4 CAPLUS
CN Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N,.alpha.-dimethyl(9CI)
(CA INDEX NAME)

NHMe CH—Me L4 ANSWER 5 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

CN Benzenemethanamine, 4-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA INDEX NAME)

MeNH-CH₂

RN 289717-61-9 CAPLUS
CN Benzenemethanamine, 5-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA INDEX NAME)

MeNH-CH₂

RN 289717-67-5 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-,
(+)- (9CI) (CA INDEX NAME)

Rotation (+).

Me NHMe C1

RN 289717-68-6 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-,
(-)- (9CI) (CA INDEX NAME)

Rotation (-).

L4 ANSWER 5 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

289717-71-1 CAPLUS Benzenemethanamine, 2-(4-chloro-3-fluorophenoxy)-5-fluoro-N-methyl- (9CI)

MeNH-CH2

289717-72-2 CAPLUS Benzenemethanamine, 2-(3-chloro-4-fluorophenoxy)-5-fluoro-N-methyl- (9CI) (CA INDEX NAME)

444888-21-5 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.,.alpha.trimethyl- (9CI) (CA INDEX NAME)

ANSWER 5 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

444888-49-7 CAPLUS Benzenemethanamine, 4-chloro-2-(3,4-dichlorophenoxy)-N-methyl-5-(methylsulfonyl) - (9CI) (CA INDEX NAME)

ANSWER 5 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

Me

444888-23-7 CAPLUS Benzenemethanamine, 4-chloro-2-(4-chlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA INDEX NAME)

MeNH-CH2

444888-24-8 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-4-methoxy-N-methyl-(9CI) (CA INDEX NAME)

менн- сн2

444888-30-6 CAPLUS Benzenemethanamine, 5-chloro-2-(3,4-dichlorophenoxy)-.alpha.-ethyl-Nmethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 6 OF 10 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2002:595509 CAPLUS DOCUMENT NUMBER: 137:135106 TITLE: Combination of a 5-HT3 receptor antagonist with a serotonin reuptake inhibitor for the treatment of

depression INVENTOR (S): Howard, Harry R. PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 20 pp. CODEN: USXXCO DOCUMENT TYPE: Patent

LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE US 2002107244 A1 20020808 US 2001-2303 20011102 EP 1230921 20020814 A1 EP 2002-250541 20020128 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR JP 2002275097 A2 20020925 JP 2002-20186 20020129

BR 2002000246 20021029 BR 2002-246 20020131 PRIORITY APPLN. INFO.: US 2001-266340P P 20010202 OTHER SOURCE(S): MARPAT 137:135106 AB The present invention relates to a method of treating depression or

anxiety in a mammal, including a human, by administering to the mammal a 5-HT3 receptor antagonist in combination with a serotonin reuptake inhibitor (SRI) antidepressant agent with improvement in sexual function and/or redn. in gastro-intestinal side effects. It also relates to pharmaceutical compns. contg. a pharmaceutically acceptable carrier, a 5-HT3 receptor antagonist and an SRI antidepressant. The ratio of the 5-HT3 receptor antagonist and the SRI antidepressant agent is between 0.001 to 1 and 1000 to 1, and esp. between 0.01 to I and 100 to 1 (no

IT 289716-94-5 289717-18-6 289717-24-4 289717-57-3 289717-58-4 289717-60-8 289717-61-9 289717-67-5 289717-68-6 289717-71-1 289717-72-2 444888-21-5 444888-23-7 444888-24-8 444888-30-6

444888-49-7 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(combination of 5-HT3 receptor antagonist with serotonin reuptake inhibitor for treatment of depression) 289716-94-5 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA

INDEX NAME)

L4 ANSWER 6 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

RN 289717-18-6 CAPLUS

N Benzenemethanamine, 4-chloro-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA

MeNH-CH₂

RN 289717-24-4 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl(9CI) (CA INDEX NAME)

NHMe CH-Me

RN 289717-57-3 CAPLUS
CN Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA INDEX NAME)

MeNH-CH₂

RN 289717-58-4 CAPLUS
CN Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N,.alpha.-dimethyl(9CI)
(CA INDEX NAME)

L4 ANSWER 6 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

NHMe CH-Me

RN 289717-60-8 CAPLUS
CN Benzenemethanamine, 4-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA INDEX NAME)

MeNH-CH₂

RN 289717-61-9 CAPLUS
CN Benzenemethanamine, 5-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA INDEX NAME)

MeNH-CH₂

Br

C1

RN 289717-67-5 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-,
(+)- (9CI) (CA INDEX NAME)

Rotation (+).

L4 ANSWER 6 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

Me_NHMe_cl

RN 289717-68-6 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-,
(-)- (9CI) (CA INDEX NAME)

Rotation (-).

Me_NHMe C1

RN 289717-71-1 CAPLUS
CN Benzenemethanamine, 2-(4-chloro-3-fluorophenoxy)-5-fluoro-N-methyl- (9CI)
(CA INDEX NAME)

MeNH-CH2

RN 289717-72-2 CAPLUS
CN Benzenemethanamine, 2-(3-chloro-4-fluorophenoxy)-5-fluoro-N-methyl- (9CI)
(CA INDEX NAME)

L4 ANSWER 6 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

MeNH—CH₂

RN 444888-21-5 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.,.alpha.trimethyl- (9CI) (CA INDEX NAME)

C1 NHMe NHMe Me

RN 444888-23-7 CAPLUS
CN Benzenemethanamine, 4-chloro-2-(4-chlorophenoxy)-5-fluoro-N-methyl- (9CI)
(CA INDEX NAME)

MeNH-CH2

RN 444888-24-8 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-4-methoxy-N-methyl(9CI) (CA INDEX NAME)

MeNH-CH₂

OMe

C1

ANSWER 6 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

444888-30-6 CAPLUS

Benzenemethanamine, 5-chloro-2-(3,4-dichlorophenoxy)-.alpha.-ethyl-Nmethyl- (9CI) (CA INDEX NAME)

RN 444888-49-7 CAPLUS Benzenemethanamine, 4-chloro-2-(3,4-dichlorophenoxy)-N-methyl-5-(methylsulfonyl) - (9CI) (CA INDEX NAME)

ANSWER 7 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued) (CH2)n(C3-C6cycloalkyl) wherein $n=0,\ 1,\ 2$ or 3; or R1 and R2 together with the nitrogen to which they are attached from an azetidine ring; Z or Y is -SR3 and the other Z or Y is halogen or -R3; wherein R3 = C1-4 alkyl optionally substituted with fluorine; except that R3 is not CF3; or Z and Y are linked so that, together with the interconnecting atoms, Z and Y form a fused 5 to 7-membered carbocyclic or heterocyclic ring, and wherein

when Z and Y form a heterocyclic ring, in addn. to carbon atoms, the linkage contains one or two heteroatoms independently selected from O, S and N; R4 and R5 independently = A-X, wherein A = -CH = CH - or - (CH2)pwhere p is 0, 1 or 2; X = H, halo, CONR6R7, SO2NR6R7, SO2NHC(=0)R6, OH, C1-4alkoxy, etc; or A-X = (un)substituted 5- or 6-membered heterocyclic ring contg. 1, 2 or 3 heteroatoms selected from N, S and O; R6 and R7 independently * H, (un) substituted alkyl; or R6 and R7 together with the

to which they are attached form a (un)substituted 4-6 membered heterocyclic ring] and there pharmaceutically acceptable salts are prepd. Thus, II was prepd. via substitution of 5-(aminosulfonyl)-2-fluoro-Nmethylbenzamide by 2,3-dihydrobenzo(b)thiophen-5-ol with successive BF3.cntdot.THF catalyzed amide redn., formylation of secondary amine, and redn. II demonstrated a serotonin re-uptake inhibition IC50 of 4.7nM. I inhibit monoamine re-uptake and in particular exhibit activity as selective serotonin reuptake inhibitors.

402910-54-7P 402910-56-9P RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(target compd.; prepn. of serotonin re-uptake inhibitors phenoxybenzylamines)

402910-54-7 CAPLUS

Benzenemethanamine, 5-bromo-2-[3-fluoro-4-(methylthio)phenoxy]-N-methyl-, hydrochloride (9CI) (CA INDEX NAME)

MeNH-CH2

● HCl

RN 402910-56-9 CAPLUS CN

Benzenemethanamine, 5-bromo-2-[3-chloro-4-(methylthio)phenoxy]-N-methyl-(9CI) (CA INDEX NAME)

L4 ANSWER 7 OF 10 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2002:171851 CAPLUS DOCUMENT NUMBER: 136:232110

Preparation of phenoxybenzylamines as selective serotonin re-uptake inhibitors INVENTOR (S): Adam, Mavis Diane; Andrews, Mark David; Elliott, Mark

Leonard; Gymer, Geoffrey Edward; Hepworth, David; Howard, Harry Ralph, Jr.; Middleton, Donald Stuart;

Stobie, Alan PATENT ASSIGNEE(S): Pfizer Limited, UK; Pfizer Inc.

SOURCE: PCT Int. Appl., 110 pp. CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

TITLE:

PATENT NO. KIND DATE APPLICATION NO. DATE ------WO 2002018333 A1 20020307 WO 2001-IB1521 20010822 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG A5 20020313 AU 2001078650 AU 2001-78650 20010822 PRIORITY APPLN. INFO.: GB 2000-21593 A 20000831

GB 2001-7116 A 20010321 WO 2001-IB1521 W 20010822

OTHER SOURCE(S): MARPAT 136:232110

$$R^{5}$$
 R^{2}
 $H_{2}N$
 Me
 Me
 Me
 Me
 Me

Title compds. I [R1 and R2 independently = H, alkyl or

ANSWER 7 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

MeNH-CH₂

REFERENCE COUNT: THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

L4 ANSWER 8 OF 10 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2001:816609 CAPLUS DOCUMENT NUMBER: 135:344284 TITLE: Process for the preparation of 2-halobenzoic acids by palladium-catalyzed regioselective halogenation of benzoic acids Kodama, Hiroki; Katsuhira, Takeshi; Nishida, Tateki; INVENTOR(S): Hino, Tomokazu; Tsubata, Kenji PATENT ASSIGNEE (S): Nihon Nohyaku Co., Ltd., Japan SOURCE: PCT Int. Appl., 29 pp. CODEN: PIXXD2 DOCUMENT TYPE: Patent LANGUAGE: Japanese FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE WO 2001083421 A1 20011108 WO 2001-JP3707 20010427 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG AU 2001052638 A5 20011112 A1 20030122 AU 2001-52638 20010427 EP 1277726 EP 2001-926032 20010427 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR JP 2002338516 A2 20021127 JP 2001-134049 20010501 PRIORITY APPLN. INFO .: JP 2000-131032 A 20000428 JP 2001-76073 A 20010316 WO 2001-JP3707 W 20010427 OTHER SOURCE(S): CASREACT 135:344284; MARPAT 135:344284 GΙ

 $(R)_n$

A process for the prepn. of 2-halobenzoic acids of the general formula [1;

A is OH, OM (wherein M is an alkali metal), or NR6R7 (wherein R6 and R7 are each H, C1-6 alkyl, optionally substituted Ph, or the like); R is H,

L4 ANSWER 9 OF 10 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2001:283913 CAPLUS DOCUMENT NUMBER:

TITLE: Preparation of biaryl ether derivatives as monoamine reuptake inhibitors

INVENTOR(S): Howard, Harry Ralph, Jr.; Adam, Mavis Diane PATENT ASSIGNEE(S): Pfizer Products Inc., USA

SOURCE: PCT Int. Appl., 52 pp.

CODEN: PIXXD2 DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE -----------. ----------WO 2001027068 20010419 WO 2000-TB1373 20000927 AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG BR 2000014733 Α 20020611 BR 2000-14733 20000927 EP 1220831 A1 20020710 EP 2000-960916 20000927 R: AT, BE, CH, DE, DK, ES, PR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL US 6410736 B1 20020625 20001019 US 2000-692335 NO 2002001659 20020408 NO 2002-1659 PRIORITY APPLN. INFO.: US 1999-159276P P 19991013 US 1999-167761P P 19991129

WO 2000-IB1373 W 20000927

OTHER SOURCE(S): MARPAT 134:310974

The title compds. [I; rings A and B can be replaced by naphthyl group; n, m = 1-3; R1, R2 = H, alkyl, alkenyl, etc.; NR1R2 = 4-8 membered satd. (un) substituted ring contg. 1-2 heteroatoms, including N atom to which R1 and R2 are attached; R3, R4 = H, alkyl optionally substituted with 1-3 F

ANSWER 8 OF 10 CAPLUS COPYRIGHT 2003 ACS C1-6 alkyl, C1-6 alkylcarbonyl, carboxyl, C1-12 alkoxycarbonyl, optionally

substituted phenylcarbonyl, or the like; n is 0 to 4; and X is Cl, Br, or iodo, or alternatively nR's may be present on benzene-constituting carbon atoms adjacent to each other and form a C3-4 alkylene or C3-4 alkenylene-fused ring.] is characterized by reacting a benzoic acid of

general formula I (X = H; A, R, n = same as above) with a halogenating agent in the presence of a Pd catalyst. This process provides regioselective halogenation under mild conditions. Thus, 0.3 g o-toluic acid, 0.5 g N-iodosuccinimide, and 11 mL DMF were added to 50 mg Pd(OAc)2 in a 50 mL glass reaction vessel and stirred at 100.degree. for 6 h to give 92% 2-iodo-6-methylbenzoic acid with 100% regioselectivity.

371771-35-6P, 2-(4-Chloro-2-methylphenoxy)-N-(1,1-dimethyl-2methylthioethyl)-6-iodobenzamide

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of 2-halobenzoic acids by regionelective halogenation of

benzoic acids in presence of palladium compd.) 371771-35-6 CAPLUS Benzamide, 2-(4-chloro-2-methylphenoxy)-N-[1,1-dimethyl-2-

(methylthio)ethyl]-6-iodo- (9CI) (CA INDEX NAME)

CH2-SMe

Me

REFERENCE COUNT:

THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

ANSWER 9 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued) atoms; CR3R4 = 4-8 membered satd. (un) substituted carbocyclic ring;

= 4-8 membered satd. (un) substituted ring contg. 1-2 heteroatoms, including N atom to which R2 is attached; X = (un) substituted Ph, heteroaryl, heterocyclyl; Y = H, halo, alkyl optionally substituted with 1-3 F atoms, etc.; Z = H, halo, alkoxy, etc.] and their pharmaceutically acceptable salts which exhibit activity as serotonin, norepinephrine, and dopamine reuptake inhibitors and can be used in the treatment of central nervous system and other disorders, were prepd. E.g., a 3-step synthesis of I [R1 = Me; R2-R4 = H; X = 5-Ph; Z = H; Y = 3,4-Cl2] was given. All exemplified compds. I showed IC50 of .ltoreq. 250 nM for serotonin reuptake inhibition, and IC50 of .ltoreq. 1000 nM for dopamine and for norepinephrine reuptake inhibition.

289717-61-9P RL: BAC (Biological activity or effector, except adverse); BSU

logical study, unclassified); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(prepn. of biaryl ether derivs. as monoamine reuptake inhibitors) 289717-61-9 CAPLUS

Benzenemethanamine, 5-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA

MeNH-CH2

INDEX NAME)

REFERENCE COUNT:

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

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L4 ANSWER 10 OF 10 CAPLUS COPYRIGHT 2003 ACS
ACCESSION NUMBER:
                        2000:608708 CAPLUS
DOCUMENT NUMBER:
                         133:207665
TITLE:
                         Preparation of phenoxybenzylamines as monoamine
                         reuptake inhibitors
INVENTOR (S):
                         Elliott, Mark Leonard; Howard, Harry Ralph, Jr.;
                         Schmidt, Christopher Joseph; Seeger, Thomas Francis
                         Pfizer Products Inc., USA
PATENT ASSIGNEE(S):
                         PCT Int. Appl., 60 pp.
SOURCE:
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT:
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PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE WO 2000050380 A1 20000831 WO 2000-IB108 20000202 W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG A1 20011121 EP 2000-900785 20000202 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO BR 2000008958 20011127 BR 2000-8958 20000202 US 2002143003 A1 20021003 US 2001-845992 20010430 NO 2001004047 20011022 Α NO 2001-4047 20010820 PRIORITY APPLN. INFO.: US 1999-121313P P 19990223 US 2000-529207 A2 20000202

W 20000202

WO 2000-IB108 OTHER SOURCE(S): MARPAT 133:207665

ROZCR3R4NR1R2 [R = (un)substituted Ph; R1,R2 = H, alk(en)yl, alkynyl; NR1R2 = heterocyclyl; R3,R4 = H or (fluoro)alkyl; R3R4 = (un)substituted alkylene; R2R3 = atoms to complete a heterocyclic ring; Z = (un) substituted phenylene] were prepd. as monoamine reuptake inhibitors (no data). Thus, 2,5-F(F3C)C6H3CHO was aroxylated by 3,4-Cl2C6H3OH and

ANSWER 10 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

289716-95-6 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N-methyl-, (2Z)-2-butenedioate (2:1) (9CI) (CA INDEX NAME) CM 1

CRN 289716-94-5 CMF C14 H12 C12 F N O

CRN 110-16-7 CMF C4 H4 O4

Double bond geometry as shown.

289717-18-6 CAPLUS Benzenemethanamine, 4-chloro-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA

ANSWER 10 OF 10 CAPLUS COPYRIGHT 2003 ACS the product reductively aminated by Me2NH to give title compd. I. 289716-89-8P 289716-93-4P 289716-94-5P 289716-95-6P 289717-18-6P 289717-19-7P 289717-23-3P 289717-24-4P 289717-25-5P 289717-26-6P 289717-28-8P 289717-35-7P 289717-36-8P 289717-57-3P 289717-58-4P 289717-60-8P 289717-61-9P 289717-67-5P 289717-68-6P 289717-71-1P 289717-72-2P RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (prepn. of phenoxybenzylamines as monoamine reuptake inhibitors) 289716-89-8 CAPLUS Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N-methyl-, hydrochloride (9CI) (CA INDEX NAME) менн- сн2

● HC1

289716-93-4 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N-methyl-, hydrochloride (9CI) (CA INDEX NAME)

● HC1

289716-94-5 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA INDEX NAME)

ANSWER 10 OF 10 CAPLUS COPYRIGHT 2003 ACS 289717-19-7 CAPLUS Benzenemethanamine, 4-chloro-2-(3,4-dichlorophenoxy)-N-methyl-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME) CRN 289717-18-6

МеNH- CH2

CMF C14 H12 C13 N O

CM 2 CRN 110-16-7 CMF C4 H4 O4

Double bond geometry as shown.

289717-23-3 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-3-fluoro-N-methyl-, (22) -2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CRN 289717-22-2 CMF C14 H12 C12 F N O

CM 2

CRN 110-16-7

ANSWER 10 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued) CMF C4 H4 04

Double bond geometry as shown.

289717-24-4 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-(9CI) (CA INDEX NAME)

NHMe

289717-25-5 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CRN 289717-24-4 CMF C15 H14 C12 F N O

CRN 110-16-7

CMF C4 H4 O4

ANSWER 10 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

RN 289717-35-7 CAPLUS Benzenemethanamine, 5-bromo-2-(3,4-dichlorophenoxy)-N-methyl-, hydrochloride (9CI) (CA INDEX NAME)

● HCl

289717-36-8 CAPLUS Benzenemethanamine, 4-bromo-2-(3,4-dichlorophenoxy)-N-methyl-, hydrochloride (9CI) (CA INDEX NAME)

менн- сн2

● HCl

289717-57-3 CAPLUS Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA

MeNH-CH2

289717-58-4 CAPLUS Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-(9CI)

L4 ANSWER 10 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued) Double bond geometry as shown.

289717-26-6 CAPLUS Benzenemethanamine, 2-(4-chlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-, hydrochloride (9CI) (CA INDEX NAME)

HC1

289717-28-8 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-4-fluoro-N-methyl-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 289717-27-7 CMF C14 H12 C12 F N O

CM 2

CRN 110-16-7

CMF C4 H4 Q4

Double bond geometry as shown.

ANSWER 10 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued) (CA INDEX NAME)

289717-60-8 CAPLUS Benzenemethanamine, 4-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA CN

289717-61-9 CAPLUS Benzenemethanamine, 5-bromo-2-(3,4-dichlorophenoxy)-N-methyl- (9CI) (CA CN INDEX NAME)

289717-67-5 CAPLUS Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N, alpha.-dimethyl-, (+) - (9CI) (CA INDEX NAME)

Rotation (+).

L4 ANSWER 10 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued)

RN 289717-68-6 CAPLUS
CN Benzenemethanamine, 2-(3,4-dichlorophenoxy)-5-fluoro-N,.alpha.-dimethyl-,
(-)- (9CI) (CA INDEX NAME)

Rotation (-).

RN 289717-71-1 CAPLUS
CN Benzenemethanamine, 2-(4-chloro-3-fluorophenoxy)-5-fluoro-N-methyl- (9CI)
(CA INDEX NAME)

MeNH-CH₂

RN 289717-72-2 CAPLUS
CN Benzenemethanamine, 2-(3-chloro-4-fluorophenoxy)-5-fluoro-N-methyl- (9CI)
(CA INDEX NAME)

L4 ANSWER 10 OF 10 CAPLUS COPYRIGHT 2003 ACS (Continued

MeNH—CH₂

CN Benzamide, 2-(3,4-dichlorophenoxy)-5-fluoro-N-methyl- (9CI) (CA INDEX NAME)

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

Kamal Saeed